

SEQUENCE LISTING

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<110> De Maria, Leonardo
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      Christensen, Lars Lehmann Hylling
      Lassen, Soren Flensted
      Ostergaard, Peter Rahbek

<120> Protease Variants

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<170> PatentIn version 3.3

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Ser Thr Thr Gly Trp His Cys Gly Thr Ile Gln Ala Arg Gly Gln Ser  
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Val Ser Tyr Pro Glu Gly Thr Val Thr Asn Met Thr Arg Thr Thr Val  
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Cys Ala Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Thr Gln  
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 Ala Glu Ala Glu Glu Leu Leu Ser Ala Gln Glu Ala Ala Ile Glu  
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Val Asp Glu Ala Ala Ala Ala Ala Ala Gly Asp Ala Tyr Gly Gly  
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Val Ser Tyr Gly Ile Glu Gly Leu Asp Glu Ile Ile Gln Asp Leu Asn  
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Ala Ala Gly Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly Arg Val  
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Thr Leu Thr Asn Leu Val Ser Arg Tyr Asn Thr Gly Gly Tyr Ala Thr  
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Val Ala Gly His Asn Gln Ala Pro Ile Gly Ser Ser Val Cys Arg Ser  
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Gly Ser Thr Thr Gly Trp His Cys Gly Thr Ile Gln Ala Arg Gly Gln  
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Ser Val Ser Tyr Pro Glu Gly Thr Val Thr Asn Met Thr Arg Thr Thr  
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Val Cys Ala Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Asn  
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Gln Ala Gln Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Arg Thr Gly  
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Leu Glu Ala Asp Glu Leu Leu Ala Ala Gln Asp Thr Ala Phe Glu	
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Val Asp Glu Ala Ala Ala Glu Ala Ala Gly Asp Ala Tyr Gly Gly	
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Ser Val Phe Asp Thr Glu Thr Leu Glu Leu Thr Val Leu Val Thr Asp	
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Ser Ala Ala Val Glu Ala Val Glu Ala Thr Gly Ala Gly Thr Glu Leu	
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gcc gcc ggt cag ccc ggg ttc gtc acc gcc ggt cac tgt ggc cgc gtg	612
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Ile Phe Pro Gly Asn Asp Ala Ala Phe Val Arg Gly Thr Ser Asn Phe	
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Thr Leu Thr Asn Leu Val Ser Arg Tyr Asn Thr Gly Gly Tyr Ala Thr	
75 80 85	
gtc gcc ggt cac aac cag gcg ccc atc ggc tcc tcc gtc tgc cgc tcc	804
Val Ala Gly His Asn Gln Ala Pro Ile Gly Ser Ser Val Cys Arg Ser	
90 95 100	
ggc tcc acc acc ggt tgg cac tgc ggc acc atc cag gcc cgc ggc cag	852
Gly Ser Thr Thr Gly Trp His Cys Gly Thr Ile Gln Ala Arg Gly Gln	
105 110 115	
tcg gtg agc tac ccc gag ggc acc gtc acc aac atg acg cgg acc acc	900
Ser Val Ser Tyr Pro Glu Gly Thr Val Thr Asn Met Thr Arg Thr Thr	
120 125 130 135	
gtg tgc gcc gag ccc ggc gac tcc ggc ggc tcc tac atc tcc ggc aac	948
Val Cys Ala Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Asn	
140 145 150	
cag gcc cag ggc gtc acc tcc ggc ggc tcc ggc aac tgc cgc acc ggc	996
Gln Ala Gln Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Arg Thr Gly	
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ggg acc acc ttc tac cag gag gtc acc ccc atg gtg aac tcc tgg ggc	1044
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Val Arg Leu Arg Thr	
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Val Asp Glu Ala Ala Ala Glu Ala Ala Gly Asp Ala Tyr Gly Gly  
-120 -115 -110

Ser Val Phe Asp Thr Glu Thr Leu Glu Leu Thr Val Leu Val Thr Asp  
-105 -100 -95 -90

Ser Ala Ala Val Glu Ala Val Glu Ala Thr Gly Ala Gly Thr Glu Leu  
-85 -80 -75

Val Ser Tyr Gly Ile Thr Gly Leu Asp Glu Ile Val Glu Glu Leu Asn  
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Ala Ala Asp Ala Val Pro Gly Val Val Gly Trp Tyr Pro Asp Val Ala  
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Gly Asp Thr Val Val Leu Glu Val Leu Glu Gly Ser Gly Ala Asp Val  
-40 -35 -30

Gly Gly Leu Leu Ala Asp Ala Gly Val Asp Ala Ser Ala Val Glu Val  
-25 -20 -15 -10

Thr Thr Thr Glu Gln Pro Glu Leu Tyr Ala Asp Ile Ile Gly Gly Leu  
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Ala Tyr Thr Met Gly Gly Arg Cys Ser Val Gly Phe Ala Ala Thr Asn  
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Ala Ala Gly Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly Arg Val  
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Gly Thr Gln Val Thr Ile Gly Asn Gly Arg Gly Val Phe Glu Gln Ser  
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Ile Phe Pro Gly Asn Asp Ala Ala Phe Val Arg Gly Thr Ser Asn Phe  
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Thr Leu Thr Asn Leu Val Ser Arg Tyr Asn Thr Gly Gly Tyr Ala Thr  
75 80 85

Val Ala Gly His Asn Gln Ala Pro Ile Gly Ser Ser Val Cys Arg Ser  
90 95 100

Gly Ser Thr Thr Gly Trp His Cys Gly Thr Ile Gln Ala Arg Gly Gln  
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Ser Val Ser Tyr Pro Glu Gly Thr Val Thr Asn Met Thr Arg Thr Thr  
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Val Cys Ala Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Asn  
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Gln Ala Gln Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Arg Thr Gly  
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Val Arg Leu Arg Thr  
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 <213> Nocardiosis alba DSM 15647 ("Protease 08")

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 -150 -145 -140  
 ccc tct cag gcc gac gag ctc ctc gag gcg cag gcc gag tcc ttc 135  
 Pro Ser Gln Ala Asp Glu Leu Leu Glu Ala Gln Ala Glu Ser Phe  
 -135 -130 -125  
 gag atc gac gag gcc gcc acc gcg gcc gca gcc gac tcc tac gcc 180  
 Glu Ile Asp Glu Ala Ala Thr Ala Ala Ala Ala Asp Ser Tyr Gly  
 -120 -115 -110  
 gcc tcc atc ttc gac acc gac agc ctc acc ctg acc gtc ctg gtc acc 228



Gly Ser Ile Phe Asp Thr Asp Ser Leu Thr Leu Thr Val Leu Val Thr	
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Asp Ala Ser Ala Val Glu Ala Val Glu Ala Ala Gly Ala Glu Ala Lys	
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Val Val Ser His Gly Met Glu Gly Leu Glu Glu Ile Val Ala Asp Leu	
-75 -70 -65 -60	
aac gcg gcc gac gct cag ccc ggc gtc gtg ggc tgg tac ccc gac atc	372
Asn Ala Ala Asp Ala Gln Pro Gly Val Val Gly Trp Tyr Pro Asp Ile	
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cac tcc gac acg gtc gtc ctc gag gtc ctc gag ggc tcc ggt gcc gac	420
His Ser Asp Thr Val Val Leu Glu Val Leu Glu Gly Ser Gly Ala Asp	
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gtg gac tcc ctg ctc gcc gac gcc ggt gtg gac acc gcc gac gtc aag	468
Val Asp Ser Leu Leu Ala Asp Ala Gly Val Asp Thr Ala Asp Val Lys	
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gtg gag agc acc acc gag cag ccc gag ctg tac gcc gac atc atc ggc	516
Val Glu Ser Thr Thr Glu Gln Pro Glu Leu Tyr Ala Asp Ile Ile Gly	
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ggt ctc gcc tac acc atg ggt ggg cgc tgc tcg gtc ggc ttc gcg gcc	564
Gly Leu Ala Tyr Thr Met Gly Gly Arg Cys Ser Val Gly Phe Ala Ala	
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acc aac gcc tcc ggc cag ccc ggg ttc gtc acc gcc ggc cac tgc ggc	612
Thr Asn Ala Ser Gly Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly	
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acc gtc ggc acc ccg gtc agc atc ggc aac ggc cag ggc gtc ttc gag	660
Thr Val Gly Thr Pro Val Ser Ile Gly Asn Gly Gln Gly Val Phe Glu	
40 45 50	
cgt tcc gtc ttc ccc ggc aac gac tcc gcc ttc gtc cgc ggc acc tcg	708
Arg Ser Val Phe Pro Gly Asn Asp Ser Ala Phe Val Arg Gly Thr Ser	
55 60 65	
aac ttc acc ctg acc aac ctg gtc agc cgc tac aac acc ggt ggt tac	756
Asn Phe Thr Leu Thr Asn Leu Val Ser Arg Tyr Asn Thr Gly Gly Tyr	
70 75 80 85	
gcg acc gtc tcc ggc tcc tcg cag gcg gcg atc ggc tcg cag atc tgc	804
Ala Thr Val Ser Gly Ser Ser Gln Ala Ala Ile Gly Ser Gln Ile Cys	
90 95 100	
cgt tcc ggc tcc acc acc ggc tgg cac tgc ggc acc gtc cag gcc cgc	852
Arg Ser Gly Ser Thr Thr Gly Trp His Cys Gly Thr Val Gln Ala Arg	
105 110 115	
ggc cag acg gtg agc tac ccc cag ggc acc gtg cag aac ctg acc cgc	900
Gly Gln Thr Val Ser Tyr Pro Gln Gly Thr Val Gln Asn Leu Thr Arg	

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Thr Asn Val Cys Ala Glu Pro Gly Asp Ser Gly Gly Ser Phe Ile Ser			
135	140	145	
ggc agc cag gcc cag ggc gtc acc tcc ggt ggc tcc ggc aac tgc tcc			996
Gly Ser Gln Ala Gln Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Ser			
150	155	160	165
ttc ggt ggc acc acc tac tac cag gag gtc aac ccg atg ctg agc agc			1044
Phe Gly Gly Thr Thr Tyr Tyr Gln Glu Val Asn Pro Met Leu Ser Ser			
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tgg ggt ctg acc ctg cgc acc tga			1068
Trp Gly Leu Thr Leu Arg Thr			
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 <213> Nocardiosis alba DSM 15647 ("Protease 08")

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Pro Ser Gln	Ala Asp Glu Leu Leu	Glu Ala Gln Ala Glu	Ser Phe
-135	-130	-125	
Glu Ile Asp	Glu Ala Ala Thr Ala	Ala Ala Ala Asp Ser	Tyr Gly
-120	-115	-110	
Gly Ser Ile	Phe Asp Thr Asp Ser	Leu Thr Leu Thr Val	Leu Val Thr
-105	-100	-95	
Asp Ala Ser	Ala Val Glu Ala Val Glu	Ala Ala Gly Ala Glu	Ala Lys
-90	-85	-80	
Val Val Ser	His Gly Met Glu Gly Leu Glu	Glu Ile Val Ala Asp	Leu
-75	-70	-65	-60
Asn Ala Ala	Asp Ala Gln Pro Gly Val	Val Gly Trp Tyr Pro	Asp Ile
-55	-50	-45	

His Ser Asp Thr Val Val Leu Glu Val Leu Glu Gly Ser Gly Ala Asp  
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Val Asp Ser Leu Leu Ala Asp Ala Gly Val Asp Thr Ala Asp Val Lys  
 -25 -20 -15

Val Glu Ser Thr Thr Glu Gln Pro Glu Leu Tyr Ala Asp Ile Ile Gly  
 -10 -5 -1 1 5

Gly Leu Ala Tyr Thr Met Gly Gly Arg Cys Ser Val Gly Phe Ala Ala  
 10 15 20

Thr Asn Ala Ser Gly Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly  
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Thr Val Gly Thr Pro Val Ser Ile Gly Asn Gly Gln Gly Val Phe Glu  
 40 45 50

Arg Ser Val Phe Pro Gly Asn Asp Ser Ala Phe Val Arg Gly Thr Ser  
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Asn Phe Thr Leu Thr Asn Leu Val Ser Arg Tyr Asn Thr Gly Gly Tyr  
 70 75 80 85

Ala Thr Val Ser Gly Ser Ser Gln Ala Ala Ile Gly Ser Gln Ile Cys  
 90 95 100

Arg Ser Gly Ser Thr Thr Gly Trp His Cys Gly Thr Val Gln Ala Arg  
 105 110 115

Gly Gln Thr Val Ser Tyr Pro Gln Gly Thr Val Gln Asn Leu Thr Arg  
 120 125 130

Thr Asn Val Cys Ala Glu Pro Gly Asp Ser Gly Gly Ser Phe Ile Ser  
 135 140 145

Gly Ser Gln Ala Gln Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Ser  
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Trp Gly Leu Thr Leu Arg Thr  
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31

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<223> Expression Cassette

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<222> (1)..(3323)  
<223> Bacillus subtilis genome sequence including  
yfmH-yfmD-yfmC-yfmB-yfmA genes

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<221> misc\_recomb  
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<223> Cat gene providing chloramphenicol resistance

<220>  
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<223> Triple PamyL-scBAN-CryIIIA promoter including mRNA stabilizing  
sequence

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<220>  
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 <223> Bacillus subtilis genome DNA including yf1S-citM genes

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Thr Ala Leu Leu Ile Ser Val Ala Phe Ser Ser Ser Ile Ala Ser	
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Ala Ala Thr Gly Ala Leu Pro Gln Ser Pro Thr Pro Glu Ala Asp	
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-150 -145 -140	
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Ser Ala Glu Ala Glu Glu Leu Leu Ala Ala Gln Asp Thr Ala Phe	
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gtc agc ggc ctg ctc gcg gac gcc ggc gtg gac gcc tcg gcc gtc gag Val Ser Gly Leu Leu Ala Asp Ala Gly Val Asp Ala Ser Ala Val Glu -25 -20 -15	6203
gtg acc acg agc gac cag ccc gag ctc tac gcc gac atc atc ggt ggt Val Thr Thr Ser Asp Gln Pro Glu Leu Tyr Ala Asp Ile Ile Gly Gly -10 -5 -1 1 5	6251
ctg gcc tac acc atg ggc ggc cgc tgt tcg gtc ggc ttc gcg gcc acc Leu Ala Tyr Thr Met Gly Gly Arg Cys Ser Val Gly Phe Ala Ala Thr 10 15 20	6299
aac gcc gcc ggt cag ccc ggg ttc gtc acc gcc ggt cac tgc ggc cgc Asn Ala Ala Gly Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly Arg 25 30 35	6347
gtg ggc acc cag gtg acc atc ggc aac ggc agg ggc gtc ttc gag cag Val Gly Thr Gln Val Thr Ile Gly Asn Gly Arg Gly Val Phe Glu Gln 40 45 50	6395
tcc gtc ttc ccc ggc aac gac gcg gcc ttc gtc cgc ggt acg tcc aac Ser Val Phe Pro Gly Asn Asp Ala Ala Phe Val Arg Gly Thr Ser Asn 55 60 65 70	6443
ttc acg ctg acc aac ctg gtc agc cgc tac aac acc ggc ggg tac gcc Phe Thr Leu Thr Asn Leu Val Ser Arg Tyr Asn Thr Gly Gly Tyr Ala 75 80 85	6491
acg gtc gcc ggt cac aac cag gcc ccc atc ggc tcc tcc gtc tgc cgc Thr Val Ala Gly His Asn Gln Ala Pro Ile Gly Ser Ser Val Cys Arg 90 95 100	6539
tcc ggc tcc acc acc ggt tgg cac tgc ggc acc atc cag gcc cgc ggc	6587

Ser	Gly	Ser	Thr	Thr	Gly	Trp	His	Cys	Gly	Thr	Ile	Gln	Ala	Arg	Gly	
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cag	tcg	gtg	agc	tac	ccc	gag	ggc	acc	gtc	acc	aac	atg	acc	cgg	acc	6635
Gln	Ser	Val	Ser	Tyr	Pro	Glu	Gly	Thr	Val	Thr	Asn	Met	Thr	Arg	Thr	
	120					125					130					
acc	gtg	tgc	gcc	gag	ccc	ggc	gac	tcc	ggc	ggc	tcc	tac	atc	tcc	ggc	6683
Thr	Val	Cys	Ala	Glu	Pro	Gly	Asp	Ser	Gly	Gly	Ser	Tyr	Ile	Ser	Gly	
	135				140					145					150	
acc	cag	gcc	cag	ggc	gtg	acc	tcc	ggc	ggc	tcc	ggc	aac	tgc	cgc	acc	6731
Thr	Gln	Ala	Gln	Gly	Val	Thr	Ser	Gly	Gly	Ser	Gly	Asn	Cys	Arg	Thr	
				155					160					165		
ggc	ggg	acc	acc	ttc	tac	cag	gag	gtc	acc	ccc	atg	gtg	aac	tcc	tgg	6779
Gly	Gly	Thr	Thr	Phe	Tyr	Gln	Glu	Val	Thr	Pro	Met	Val	Asn	Ser	Trp	
			170					175					180			
ggc	gtc	cgt	ctc	cgg	acc	taatcgc	atg	ttcaatccgc	tccataatcg							6827
Gly	Val	Arg	Leu	Arg	Thr											
	185															
gtc	gac	gcg	cgg	ttc	gcg	ctc	gac	agca	cat	cacc	gaa	atattat	gga	agaaa	atattc	6887
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gct	ctt	caag	ttat	cctttt	agct	atg	cat	ggg	gaat	cgg	aaagt	catct	aaa	atct	atg	7187
cccaaa	acaa	tgt	catt	gac	gtac	cggg	ac	tgt	cag	ctgc	taaa	acgatc	agc	gtatt	tca	7247
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aag	aa	agtga	aaa	acacaaa	gggt	gcta	acc	cttt	gtgttt	ttta	atta	aat	taaa	atgttt		7487
atta	act	tag	tta	agg	agta	gaat	ggaaaa	gggg	atc	gga	aaaca	agtat	atag	gagg	gag	7547
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aac	ctaa	agc	ttgg	cattt	g	tttgc	gattt	ttgt	cgcaac	aatt	atc	ggc	ttt	atct	cca	7727
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cact	at	caat	tgag	gata	ca	tta	agcggat	tcg	ggaataa	gacc	attt	gg	ctt	atc	gtta	7847

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tgtctgttga ttttctgcaa acggagacag agcaaagcat caaaaagcat ttgagaaatt 9647  
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 aaagcatcag aaaggatacg ctcgggcttg aaccgcatga gatcgcggt ctatatcgtg 9767  
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 <211> 380  
 <212> PRT  
 <213> Artificial

<220>  
 <223> Synthetic Construct

<400> 14

Met Lys Lys Pro Leu Gly Lys Ile Val Ala Ser Thr Ala Leu Leu  
 -190 -185 -180

Ile Ser Val Ala Phe Ser Ser Ser Ile Ala Ser Ala Ala Thr Gly  
 -175 -170 -165

Ala Leu Pro Gln Ser Pro Thr Pro Glu Ala Asp Ala Val Ser Met  
 -160 -155 -150

Gln Glu Ala Leu Gln Arg Asp Leu Asp Leu Thr Ser Ala Glu Ala  
 -145 -140 -135

Glu Glu Leu Leu Ala Ala Gln Asp Thr Ala Phe Glu Val Asp Glu  
 -130 -125 -120

Ala Ala Ala Glu Ala Ala Gly Asp Ala Tyr Gly Gly Ser Val Phe  
 -115 -110 -105

Asp Thr Glu Ser Leu Glu Leu Thr Val Leu Val Thr Asp Ala Ala Ala

-100					-95					-90					
Val	Glu	Ala	Val	Glu	Ala	Thr	Gly	Ala	Gly	Thr	Glu	Leu	Val	Ser	Tyr
-85						-80					-75				
Gly	Ile	Asp	Gly	Leu	Asp	Glu	Ile	Val	Gln	Glu	Leu	Asn	Ala	Ala	Asp
-70					-65					-60					-55
Ala	Val	Pro	Gly	Val	Val	Gly	Trp	Tyr	Pro	Asp	Val	Ala	Gly	Asp	Thr
				-50					-45					-40	
Val	Val	Leu	Glu	Val	Leu	Glu	Gly	Ser	Gly	Ala	Asp	Val	Ser	Gly	Leu
			-35					-30					-25		
Leu	Ala	Asp	Ala	Gly	Val	Asp	Ala	Ser	Ala	Val	Glu	Val	Thr	Thr	Ser
		-20					-15					-10			
Asp	Gln	Pro	Glu	Leu	Tyr	Ala	Asp	Ile	Ile	Gly	Gly	Leu	Ala	Tyr	Thr
-5					-1	1				5					10
Met	Gly	Gly	Arg	Cys	Ser	Val	Gly	Phe	Ala	Ala	Thr	Asn	Ala	Ala	Gly
				15					20					25	
Gln	Pro	Gly	Phe	Val	Thr	Ala	Gly	His	Cys	Gly	Arg	Val	Gly	Thr	Gln
			30					35					40		
Val	Thr	Ile	Gly	Asn	Gly	Arg	Gly	Val	Phe	Glu	Gln	Ser	Val	Phe	Pro
		45					50					55			
Gly	Asn	Asp	Ala	Ala	Phe	Val	Arg	Gly	Thr	Ser	Asn	Phe	Thr	Leu	Thr
	60						65				70				
Asn	Leu	Val	Ser	Arg	Tyr	Asn	Thr	Gly	Gly	Tyr	Ala	Thr	Val	Ala	Gly
75					80					85					90
His	Asn	Gln	Ala	Pro	Ile	Gly	Ser	Ser	Val	Cys	Arg	Ser	Gly	Ser	Thr
				95					100					105	
Thr	Gly	Trp	His	Cys	Gly	Thr	Ile	Gln	Ala	Arg	Gly	Gln	Ser	Val	Ser
			110					115					120		
Tyr	Pro	Glu	Gly	Thr	Val	Thr	Asn	Met	Thr	Arg	Thr	Thr	Val	Cys	Ala
		125					130					135			

Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Thr Gln Ala Gln  
 140 145 150

Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Arg Thr Gly Gly Thr Thr  
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Phe Tyr Gln Glu Val Thr Pro Met Val Asn Ser Trp Gly Val Arg Leu  
 175 180 185

Arg Thr

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 <211> 35  
 <212> DNA  
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<220>  
 <223> Primer

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 ggagctctga aaaaaaggag aggataaaga atgaa 35

<210> 16  
 <211> 29  
 <212> DNA  
 <213> Artificial

<220>  
 <223> Primer

<400> 16  
 gcgttcgat aatcgcggtg acaatgccg 29

<210> 17  
 <211> 29  
 <212> DNA  
 <213> Artificial

<220>  
 <223> Primer

<400> 17  
 ttcattgagtc tgcgccctga gatcctctg 29

<210> 18  
 <211> 30  
 <212> DNA

<213> Artificial  
 <220>  
 <223> Primer  
 <400> 18  
 taatcgcatg ttcaatccgc tccataatcg 30

<210> 19  
 <211> 44  
 <212> DNA  
 <213> Artificial  
 <220>  
 <223> Primer  
 <400> 19  
 cccaacggtt tcttcattct ttatcctctc ctttttttca gage 44

<210> 20  
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 <213> Artificial  
 <220>  
 <223> Protease 22

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 <220>  
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 <222> (1)..(81)  
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 <222> (577)..(1164)

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 Met Lys Lys Pro Leu Gly Lys Ile Val Ala Ser Thr Ala Leu Leu  
 -190 -185 -180

att tca gtg gca ttt agc tca tct att gca tca gca gct aca gga 90  
 Ile Ser Val Ala Phe Ser Ser Ser Ile Ala Ser Ala Ala Thr Gly  
 -175 -170 -165

gca tta ccg cag tct ccg aca ccg gaa gca gat gca gtc tca atg 135  
 Ala Leu Pro Gln Ser Pro Thr Pro Glu Ala Asp Ala Val Ser Met  
 -160 -155 -150

caa gaa gca ctg caa aga gat ctt gat ctt aca tca gca gaa gca 180  
 Gln Glu Ala Leu Gln Arg Asp Leu Asp Leu Thr Ser Ala Glu Ala

-145	-140	-135	
gaa gaa ctt ctt gct gca caa gat aca gca ttt gaa gtg gat gaa			225
Glu Glu Leu Leu Ala Ala Gln Asp Thr Ala Phe Glu Val Asp Glu			
-130	-125	-120	
gca gcg gca gaa gca gca gga gat gca tat ggc ggc tca gtt ttt			270
Ala Ala Ala Glu Ala Ala Gly Asp Ala Tyr Gly Gly Ser Val Phe			
-115	-110	-105	
gat aca gaa tca ctt gaa ctt aca gtt ctt gtt aca gat gca gca gca			318
Asp Thr Glu Ser Leu Glu Leu Thr Val Leu Val Thr Asp Ala Ala Ala			
-100	-95	-90	
gtt gaa gca gtt gaa gca aca gga gca gga aca gta ctt gtt tca tat			366
Val Glu Ala Val Glu Ala Thr Gly Ala Gly Thr Val Leu Val Ser Tyr			
-85	-80	-75	
gga att gat ggc ctt gat gaa att gtt caa gaa ctg aat gca gct gat			414
Gly Ile Asp Gly Leu Asp Glu Ile Val Gln Glu Leu Asn Ala Ala Asp			
-70	-65	-60	-55
gct gtt ccg ggc gtt gtt ggc tgg tat ccg gat gtt gct gga gat aca			462
Ala Val Pro Gly Val Val Gly Trp Tyr Pro Asp Val Ala Gly Asp Thr			
-50	-45	-40	
gtt gtc ctt gaa gtt ctt gaa gga tca ggc gca gat gtt tca ggc ctg			510
Val Val Leu Glu Val Leu Glu Gly Ser Gly Ala Asp Val Ser Gly Leu			
-35	-30	-25	
ctg gca gac gca gga gtc gat gca tca gca gtt gaa gtt aca aca tca			558
Leu Ala Asp Ala Gly Val Asp Ala Ser Ala Val Glu Val Thr Thr Ser			
-20	-15	-10	
gat caa ccg gaa ctt tat gca gat att att ggc ggc ctg gca tat tat			606
Asp Gln Pro Glu Leu Tyr Ala Asp Ile Ile Gly Gly Leu Ala Tyr Tyr			
-5	-1 1	5	10
atg ggc ggc aga tgc agc gtt ggc ttt gca gca aca aat gca tca ggc			654
Met Gly Gly Arg Cys Ser Val Gly Phe Ala Ala Thr Asn Ala Ser Gly			
15	20	25	
caa ccg ggc ttt gtt aca gca ggc cat tgc ggc aca gtt ggc aca cca			702
Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly Thr Val Gly Thr Pro			
30	35	40	
gtt tca att ggc aat ggc aaa ggc gtt ttt gaa cga agc att ttt ccg			750
Val Ser Ile Gly Asn Gly Lys Gly Val Phe Glu Arg Ser Ile Phe Pro			
45	50	55	
ggc aat gat tca gca ttt gtt aga ggc aca tca aat ttt aca ctt aca			798
Gly Asn Asp Ser Ala Phe Val Arg Gly Thr Ser Asn Phe Thr Leu Thr			
60	65	70	
aat ctg gtt tca aga tat aat tca ggc ggc tat gca aca gtt gca ggc			846
Asn Leu Val Ser Arg Tyr Asn Ser Gly Gly Tyr Ala Thr Val Ala Gly			
75	80	85	90



cat aat caa gca ccg att ggc tca gca gtt tgc aga tca ggc tca aca	894
His Asn Gln Ala Pro Ile Gly Ser Ala Val Cys Arg Ser Gly Ser Thr	
95 100 105	
aca ggc tgg cat tgc ggc aca att caa gca aga aat caa aca gtt agg	942
Thr Gly Trp His Cys Gly Thr Ile Gln Ala Arg Asn Gln Thr Val Arg	
110 115 120	
tat ccg caa ggc aca gtt tat agt ctg aca aga aca aca gtt tgt gca	990
Tyr Pro Gln Gly Thr Val Tyr Ser Leu Thr Arg Thr Thr Val Cys Ala	
125 130 135	
gaa ccg ggc gat tca ggc ggc tca tat att agc ggc act caa gca caa	1038
Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Thr Gln Ala Gln	
140 145 150	
ggc gtt aca tca ggc ggc tca ggc aat tgc agt gct ggc ggc aca aca	1086
Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Ser Ala Gly Gly Thr Thr	
155 160 165 170	
tat tac caa gaa gtt aat ccg atg ctt agt tca tgg ggc ctt aca ctt	1134
Tyr Tyr Gln Glu Val Asn Pro Met Leu Ser Ser Trp Gly Leu Thr Leu	
175 180 185	
aga aca caa tcg cat gtt caa tcc gct cca	1164
Arg Thr Gln Ser His Val Gln Ser Ala Pro	
190 195	

<210> 21  
 <211> 388  
 <212> PRT  
 <213> Artificial

<220>  
 <223> Synthetic Construct

<400> 21

Met Lys Lys Pro Leu Gly Lys Ile Val Ala Ser Thr Ala Leu Leu	
-190 -185 -180	
Ile Ser Val Ala Phe Ser Ser Ser Ile Ala Ser Ala Ala Thr Gly	
-175 -170 -165	
Ala Leu Pro Gln Ser Pro Thr Pro Glu Ala Asp Ala Val Ser Met	
-160 -155 -150	
Gln Glu Ala Leu Gln Arg Asp Leu Asp Leu Thr Ser Ala Glu Ala	
-145 -140 -135	
Glu Glu Leu Leu Ala Ala Gln Asp Thr Ala Phe Glu Val Asp Glu	

-130		-125		-120
Ala Ala Ala Glu Ala Ala Gly Asp Ala Tyr Gly Gly Ser Val Phe				
-115		-110		-105
Asp Thr Glu Ser Leu Glu Leu Thr Val Leu Val Thr Asp Ala Ala Ala				
-100		-95		-90
Val Glu Ala Val Glu Ala Thr Gly Ala Gly Thr Val Leu Val Ser Tyr				
-85		-80		-75
Gly Ile Asp Gly Leu Asp Glu Ile Val Gln Glu Leu Asn Ala Ala Asp				
-70		-65		-60
				-55
Ala Val Pro Gly Val Val Gly Trp Tyr Pro Asp Val Ala Gly Asp Thr				
	-50		-45	-40
Val Val Leu Glu Val Leu Glu Gly Ser Gly Ala Asp Val Ser Gly Leu				
	-35		-30	-25
Leu Ala Asp Ala Gly Val Asp Ala Ser Ala Val Glu Val Thr Thr Ser				
	-20		-15	-10
Asp Gln Pro Glu Leu Tyr Ala Asp Ile Ile Gly Gly Leu Ala Tyr Tyr				
-5		-1	1	5
				10
Met Gly Gly Arg Cys Ser Val Gly Phe Ala Ala Thr Asn Ala Ser Gly				
	15		20	25
Gln Pro Gly Phe Val Thr Ala Gly His Cys Gly Thr Val Gly Thr Pro				
	30		35	40
Val Ser Ile Gly Asn Gly Lys Gly Val Phe Glu Arg Ser Ile Phe Pro				
	45		50	55
Gly Asn Asp Ser Ala Phe Val Arg Gly Thr Ser Asn Phe Thr Leu Thr				
	60		65	70
Asn Leu Val Ser Arg Tyr Asn Ser Gly Gly Tyr Ala Thr Val Ala Gly				
75		80		85
				90
His Asn Gln Ala Pro Ile Gly Ser Ala Val Cys Arg Ser Gly Ser Thr				
	95		100	105

Thr Gly Trp His Cys Gly Thr Ile Gln Ala Arg Asn Gln Thr Val Arg  
110 115 120

Tyr Pro Gln Gly Thr Val Tyr Ser Leu Thr Arg Thr Thr Val Cys Ala  
125 130 135

Glu Pro Gly Asp Ser Gly Gly Ser Tyr Ile Ser Gly Thr Gln Ala Gln  
140 145 150

Gly Val Thr Ser Gly Gly Ser Gly Asn Cys Ser Ala Gly Gly Thr Thr  
155 160 165 170

Tyr Tyr Gln Glu Val Asn Pro Met Leu Ser Ser Trp Gly Leu Thr Leu  
175 180 185

Arg Thr Gln Ser His Val Gln Ser Ala Pro  
190 195